

## WEST Search History

DATE: Thursday, May 12, 2005

<u>Hide?</u>	<u>Set Name</u>	<u>Query</u>	<u>Hit Count</u>
<i>DB=USPT; PLUR=YES; OP=AND</i>			
<input type="checkbox"/>	L1	heavychain or heavy-chain or hc or h-c or carboxyterminal or carboxylterminal or carboxy-terminal or carboxyl-terminal or (receptor near3 binding) or (targeting near3 moiety) or (receptor near3 domain) or (binding near3 moiety) or (receptor near3 moiety) or rbonthc or rbont-ch or bonthc or bont-hc	54884
<i>DB=PGPB,USPT,USOC,EPAB,JPAB,DWPI,TDBD; PLUR=YES; OP=AND</i>			
<input type="checkbox"/>	L2	heavychain or heavy-chain or hc or h-c or carboxyterminal or carboxylterminal or carboxy-terminal or carboxyl-terminal or (receptor near3 binding) or (targeting near3 moiety) or (receptor near3 domain) or (binding near3 moiety) or (receptor near3 moiety) or rbonthc or rbont-ch or bonthc or bont-hc	152450
<input type="checkbox"/>	L3	botulin or botulinum or botinolysin or botulism or botox or btx or neurotoxin or neuro-toxin	11128
<input type="checkbox"/>	L4	L3 and (l1 or l2)	3434
<input type="checkbox"/>	L5	L4 and clostrid\$	1416
<input type="checkbox"/>	L6	(l1 or l2).clm. and l3.clm.	31

END OF SEARCH HISTORY

---

- 1. 20050020519. 10 Sep 04. 27 Jan 05. Modulation of insulin-regulated aminopeptidase (irap)/angiotensin iv (at4) receptor activity. Albiston, Anthony L., et al. 514/44; 435/6 435/7.2 514/12 A61K048/00 C12Q001/68 G01N033/53 G01N033/567.
- 2. 20040265935. 09 Apr 04. 30 Dec 04. Botulinum toxin a peptides and methods of predicting and reducing immunoresistance to botulinum toxin therapy. Atassi, M. Zouhair. 435/7.32; G01N033/554 G01N033/569.
- 3. 20040180048. 14 Jul 03. 16 Sep 04. Neuronal and retinal gene expression patterns. Zack, Donald Jeffery, et al. 424/143.1; 435/368 A61K039/395 C12N005/08.
- 4. 20040147716. 02 Jul 03. 29 Jul 04. Peptides comprising aromatic D-amino acids and methods of use. Anderson, Byron E.. 530/329; 530/330 C07K007/06.
- 5. 20040142455. 05 Dec 03. 22 Jul 04. Recombinant botulinum toxins having a soluble C-terminal portion of a heavy chain, an N-terminal portion of a heavy chain and a light chain. Williams, James A.. 435/252.33; 435/254.2 435/320.1 435/348 435/69.3 530/350 536/23.7 C12P021/02 C12N001/21 C12N001/18 C12N005/06.
- 6. 20040115215. 05 Dec 03. 17 Jun 04. Recombinant botulinum toxins with a soluble C-terminal portion, an N-terminal portion and a light chain. Williams, James A.. 424/184.1; A61K039/395 A61K039/00 A61K039/38.
- 7. 20040110284. 04 Sep 03. 10 Jun 04. Antibodies against type a botulinum neurotoxin. Bavari, Sina, et al. 435/326; 530/388.1 C12Q001/68 C12N005/06 C07K016/40.
- 8. 20040039212. 25 Aug 03. 26 Feb 04. Sphingolipid derivatives and their methods of use. Liotta, Dennis C., et al. 548/566; 549/491 549/74 554/36 C11C003/00 C07D333/12 C07D207/08.
- 9. 20040018589. 25 Jul 02. 29 Jan 04. Method for producing biologically active botulinum neurotoxins through recombinant DNA technique. Zhong, Jun. 435/69.1; 435/252.3 C12P021/02 C12N001/21.
- 10. 20040013687. 02 Jun 03. 22 Jan 04. Compositions and methods for transepithelial molecular transport. Simpson, Lance, et al. 424/190.1; A61K039/02.
- 11. 20030232827. 10 Feb 03. 18 Dec 03. Therapeutic compounds. Meltzer, Peter C., et al. 514/232.8; 514/253.04 514/254.11 514/304 514/320 514/434 514/469 544/127 544/145 544/150 . 544/362 546/134 546/135 546/196 546/202 549/23 549/397 A61K031/5377 A61K031/496 A61K031/46 A61K031/452 A61K031/453 A61K031/382 A61K031/353 C07D413/02 C07D49/02.
- 12. 20030221201. 04 Mar 03. 27 Nov 03. Modified transferrin fusion proteins. Prior, Christopher P., et al. 800/7; 424/85.5 514/6 530/350 530/351 530/400 A61K038/21 A61K038/40 A01K067/027 C07K014/79.
- 13. 20030219457. 15 Oct 02. 27 Nov 03. Soluble recombinant botulinum toxins. Williams, James A.. 424/199.1; 424/186.1 424/234.1 435/6 C12Q001/68 A61K039/12 A61K039/02.
- 14. 20030215468. 30 Jan 03. 20 Nov 03. Soluble recombinant botulinum toxin proteins. Williams, James A., et al. 424/239.1; 435/252.3 435/70.21 530/388.4 A61K039/08 C12P021/04 C12N001/21

---

C07K016/12.

- 15. 20030166238. 12 Sep 02. 04 Sep 03. Recombinant toxin fragments. Shone, Clifford Charles, et al. 435/219; 435/252.3 435/69.7 530/350 536/23.7 C12N009/50 C07H021/04 C12P021/04 C12N001/21 C07K014/33.
- 16. 20030088074. 28 May 02. 08 May 03. Recombinant bivalent monospecific immunoglobulin having at least two variable fragments of heavy chains of an immunoglobulin devoid of light chains. Hamers, Raymond, et al. 530/387.1; C07K016/00.
- 17. 20030059912. 27 Aug 02. 27 Mar 03. Hybrid protein for inhibiting the degranulation of mastocytes and the use thereof. Bigalke, Hans, et al. 435/188.5; 424/178.1 435/219 C12N009/50 C07K016/40 A61K039/40.
- 18. 20030009025. 20 Jul 01. 09 Jan 03. Recombinant vaccine against botulinum neurotoxin. Smith, Leonard A., et al. 536/23.7; 435/252.3 435/254.23 435/320.1 435/69.1 C07H021/04 C12P021/02 C12N001/21 C12N001/18.
- 19. 20020137886. 29 Nov 00. 26 Sep 02. Neurotoxins with enhanced target specificity. Lin, Wei-Jen, et al. 530/350; 435/325 435/69.1 536/23.7 C07K014/24 C07H021/04 C12P021/02.
- 20. 20020068699. 23 Aug 01. 06 Jun 02. Clostridial toxin derivatives and methods for treating pain. Donovan, Stephen. 514/12; 530/350 A61K039/08 C07K014/33.
- 21. 20020044950. 23 Feb 99. 18 Apr 02. RECOMBINANT TOXIN FRAGMENTS. SHONE, CLIFFORD CHARLES, et al. 424/236.1; 424/238.1 424/239.1 A61K039/02 A61K039/05 A61K039/08.
- 22. 6822076. 27 Aug 02; 23 Nov 04. Hybrid protein for inhibiting the degranulation of mastocytes and the use thereof. Bigalke; Hans, et al. 530/350; 424/192.1 435/7.1 530/300. C07K001/00.
- 23. 6713444. 12 May 00; 30 Mar 04. Buforin I as a specific inhibitor and therapeutic agent for botulinum toxin B and tetanus neurotoxins. Garcia; Gregory E., et al. 514/2; 424/239.1 424/9.1 435/252.7 514/13 514/21 530/324 530/326 530/333 530/344. A61K038/00 C07K014/00.
- 24. 6632440. 29 May 01; 14 Oct 03. Methods and compounds for the treatment of mucus hypersecretion. Quinn; Conrad Padraig, et al. 424/239.1; 424/236.1 424/282.1 424/434 424/810 435/325 435/368 435/371 435/6 435/69.1 435/7.1 514/12 514/14 514/2 530/350. A61K039/68 A61K039/00 C07K014/00.
- 25. 6573244. 12 May 00; 03 Jun 03. Previns as specific inhibitors and therapeutic agents for Botulinum toxin B and Tetanus neurotoxins. Gordon; Richard K., et al. 514/15; 424/185.1 435/69.1 536/23.1 536/23.4. A61K038/00 C07H021/02.
- 26. 6461617. 23 Feb 99; 08 Oct 02. Recombinant toxin fragments. Shone; Clifford Charles, et al. 424/236.1; 424/157.1 424/164.1 424/167.1 424/178.1 424/179.1 424/184.1 424/234.1 424/235.1 424/239.1 424/247.1 435/252.33 435/69.1 435/69.7 435/70.1 435/71.1 435/71.2 530/300 530/350 530/825 536/23.4 536/23.7. A61K039/02 A61K039/38 A61K039/00 C12P021/06 C12P021/04.
- 27. 5939070. 28 Oct 96; 17 Aug 99. Hybrid botulinal neurotoxins. Johnson; Eric A., et al. 424/194.1; 424/239.1 435/220 435/842 514/12 530/350 530/402 530/412 530/825 536/23.2 536/23.7.

A61K039/385 A61K039/08 C12P021/06 C12N009/52.

28. 5770572. 30 Jun 89; 23 Jun 98. Methods and compositions using molecular decoyants for ameliorating the undesired effects of foreign agents which bind to endogenous receptors. Gershoni; Jonathan M.. 514/13; 514/2 514/21 530/326 530/350. A61K038/00 A61K038/02 A61K038/04.

29. 5401243. 12 Apr 93; 28 Mar 95. Controlled administration of chemodenervating pharmaceuticals. Borodic; Gary E.. 604/511; 128/898. A61M031/00.

30. 5387503. 12 Nov 92; 07 Feb 95. Assay method using internal calibration to measure the amount of analyte in a sample. Selmer; Johan, et al. 435/5; 435/7.2 435/7.21 435/7.23 435/7.32 435/7.4 435/7.8 435/7.94 435/7.95 435/967 435/975 436/501 436/518 436/807. G01N033/543 G01N033/566 G01N033/569.

31. 5183462. 21 Aug 90; 02 Feb 93. Controlled administration of chemodenervating pharmaceuticals. Borodic; Gary E.. 604/506; 128/898. A61M031/00.

[Generate Collection](#)

[Print](#)

Terms	Documents
(L1 or L2).clm. and L3.clm.	31

[Prev Page](#)   [Next Page](#)   [Go to Doc#](#)

US-PAT-NO: 6713444

DOCUMENT-IDENTIFIER: US 6713444 B1

TITLE: Buforin I as a specific inhibitor and therapeutic agent for botulinum toxin B and tetanus neurotoxins

DATE-ISSUED: March 30, 2004

INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Garcia; Gregory E.	Germantown	MD		
Gordon; Richard K.	Potomac	MD		
Moorad; Debbie R.	Rockville	MD		
Doctor; Bhupendra P.	Potomac	MD		

ASSIGNEE-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY	TYPE CODE
The United States of America as represented by the Secretary of the Army	Washington DC				06

APPL-NO: 09/ 570023 [PALM]

DATE FILED: May 12, 2000

PARENT-CASE:

CROSS REFERENCE TO RELATED APPLICATIONS This application is based on provisional application No. 60/134,216 filed May 14, 1999.

INT-CL: [07] A61 K 38/00, C07 K 14/00

US-CL-ISSUED: 514/2; 514/13, 514/21, 530/324, 530/326, 530/333, 530/344, 424/239.1, 424/9.1, 435/252.7

US-CL-CURRENT: 514/2; 424/239.1, 424/9.1, 435/252.7, 514/13, 514/21, 530/324, 530/326, 530/333, 530/344

FIELD-OF-SEARCH: 514/12, 514/13, 514/21, 530/324, 530/326, 530/333, 530/344, 424/239.1, 424/9.1, 435/252.7

PRIOR-ART-DISCLOSED:

U.S. PATENT DOCUMENTS

Search Selected |  Search All |  Clear

PAT-NO	ISSUE-DATE	PATENTEE-NAME	US-CL
<u>5936063</u>	August 1999	Kim et al.	530/324

6573244

June 2003

Gordon et al.

514/15

## OTHER PUBLICATIONS

Park et al., A novel antimicrobial peptide form bufo bufo gargarizans. Biochem. Biophys. Res. Commun. 218, 408-413 (1996).\*

Garcia et al., Botulinum B toxin activity is inhibited by Buforin I. FASEB Journal vol. 12, No. 8, pp A1472. (Apr. 1998).

ART-UNIT: 1653

PRIMARY-EXAMINER: Low; Christopher S. F.

ASSISTANT-EXAMINER: Kam; Chih-Min

ATTY-AGENT-FIRM: Arwine; Elizabeth

ABSTRACT:

The compounds of the invention are generally described by the formula:

X.sub.1 X.sub.2 B.sub.3 X.sub.4 B.sub.5 X\*.sub.6 X.sub.7 X.sub.8

B.sub.9 X.sub.10 B.sub.11 X.sub.12 B.sub.13 X.sub.14

B.sub.15 X.sub.16 B.sub.17 X\*.sub.18 X\*.sub.19 B.sub.20

X.sub.21 X.sub.22 X.sub.23 Q.sub.24 F.sub.25 Z\*.sub.26 X.sub.27

X.sub.28 B.sub.29 X.sub.30 B.sub.31 B.sub.32 X.sub.33 X.sub.34

B.sub.35 B.sub.36 X.sub.37 Z.sub.38 Z.sub.39 (1)

and the salts, esters, amides, and acyl forms thereof. Up to 15 amino acids may be truncated from the N-terminus and up to 6 amino acids may be truncated from the C-terminus. Each position represented by a letter indicates a single amino acid residue wherein B is a basic or polar/large amino acid or a modified form thereof; X is a small or hydrophobic amino acid or a modified form thereof; X\* is a small or polar/large amino acid or a modified form thereof; Z is a polar/large or hydrophobic amino acid or a modified form thereof; Z\* is Proline or a polar/large or hydrophobic amino acid or a modified form thereof. These compounds may be used to inhibit the protease activity of Botulinum B and tetanus toxins.

12 Claims, 10 Drawing figures

PGPUB-DOCUMENT-NUMBER: 20020068699

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20020068699 A1

TITLE: Clostridial toxin derivatives and methods for treating pain

PUBLICATION-DATE: June 6, 2002

INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
Donovan, Stephen	Capistrano Beach	CA	US	

ASSIGNEE-INFORMATION:

NAME	CITY	STATE	COUNTRY	TYPE CODE
Allergan Sales, Inc.	Irvine	CA	US	02

APPL-NO: 09/ 938112 [PALM]

DATE FILED: August 23, 2001

RELATED-US-APPL-DATA:

Application 09/938112 is a division-of US application 09/489667, filed January 19, 2000, PENDING

INT-CL: [07] A61 K 39/08, C07 K 14/33

US-CL-PUBLISHED: 514/12; 530/350

US-CL-CURRENT: 514/12; 530/350

ABSTRACT:

Agents for treating pain, methods for producing the agents and methods for treating pain by administration to a patient of a therapeutically effective amount of the agent. The agent can include a clostridial neurotoxin, or a component or fragment or derivative thereof, attached to a targeting moiety, wherein the targeting moiety is selected from a group consisting of transmission compounds which can be released from neurons upon the transmission of pain signals by the neurons, and compounds substantially similar to the transmission compounds.

DOCUMENT-IDENTIFIER: US 20040147716 A1

TITLE: Peptides comprising aromatic D-amino acids and methods of use

CLAIMS:

14. The method of claim 12, wherein the toxin is selected from the group consisting of botulinum toxins, ricin toxins, cholera toxins, and anthrax toxins or toxin subcomponents.

31. A method of reducing the ConA lectin binding to at least one of its receptors comprising delivering to the mammal a D-peptide comprising a pentapeptide core selected from the group consisting of Xaa.sub.1YYFF and Xaa.sub.1FYFF wherein Xaa.sub.1 is an amino acid of the D- or L-configuration independently selected from the group consisting of D, E, K, R, H, N, Q, C, S, T, G, A, V, L, I, M and P.

39. A method of reducing binding of TNFA to a TNFA receptor comprising delivering to the mammal a D-peptide comprising a pentapeptide core selected from the group consisting of FFFXaa.sub.1F, YFXaa.sub.1FF, YFYFXaa.sub.1, YWXaa.sub.1FF, WXaa.sub.1YXaa.sub.2F, WXaa.sub.1YFXaa.sub.2 and WXaa.sub.1FFXaa.sub.2 wherein Xaa.sub.1 and Xaa.sub.2 are amino acids of the D- or L-configuration independently selected from the group consisting of D, E, K, R, H, N, Q, C, S, T, G, A, V, L, I, M and P.

41. A method of reducing the binding of TGF.beta.1 to a TNF.beta.1 receptor comprising delivering to the mammal a D-peptide comprising a pentapeptide core selected from the group consisting of FFFXaa.sub.1W, FWFXaa.sub.1Xaa.sub.2, FYXaa.sub.1YF, FWXaa.sub.1Xaa.sub.2Xaa.sub.3, FXaa.sub.1YYW, FXaa.sub.1YYXaa.sub.2, FWXaa.sub.1WY, FFWYW, FXaa.sub.1Xaa.sub.2FXaa.sub.3, FYWXaa.sub.1Y, FYWXaa.sub.1W, FXaa.sub.1YFXaa.sub.2, FYYYYXaa.sub.1, FWXaa.sub.1FF and FFXaa.sub.1WW wherein Xaa.sub.1, Xaa.sub.2 and Xaa.sub.3 are amino acids of the D- or L-configuration independently selected from the group consisting of D, E, K, R, H, N, Q, C, S, T, G, A, V, L, I, M and P.

PGPUB-DOCUMENT-NUMBER: 20040265935

PGPUB-FILING-TYPE: new

DOCUMENT-IDENTIFIER: US 20040265935 A1

TITLE: Botulinum toxin a peptides and methods of predicting and reducing immunoresistance to botulinum toxin therapy

PUBLICATION-DATE: December 30, 2004

INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
Atassi, M. Zouhair	Houston	TX	US	

APPL-NO: 10/ 821669 [PALM]

DATE FILED: April 9, 2004

RELATED-US-APPL-DATA:

Application is a non-provisional-of-provisional application 60/462754, filed April 11, 2003,

INT-CL: [07] G01 N 33/554, G01 N 33/569

US-CL-PUBLISHED: 435/007.32

US-CL-CURRENT: 435/7.32

REPRESENTATIVE-FIGURES: NONE

ABSTRACT:

The present invention provides BoNT/A peptides as well as methods of predicting or determining immunoresistance to botulinum toxin therapy in an individual using BoNT/A peptides.

[0001] This application is based on, and claims the benefit of, U.S. Provisional Application No. 60/462,754, filed Apr. 11, 2003, and entitled Botulinum Toxin A Peptides And Methods Of Predicting And Reducing Immunoresistance To Botulinum Toxin Therapy, and which is incorporated herein by reference.

DOCUMENT-IDENTIFIER: US 20040265935 A1

TITLE: Botulinum toxin a peptides and methods of predicting and reducing immunoresistance to botulinum toxin therapy

CLAIMS:

1. A method of predicting or determining immunoresistance to botulinum toxin therapy in an individual, comprising determining the presence or absence in said individual of antibodies immunoreactive with two or more amino acid sequences selected from the group: 785-803 of SEQ ID NO: 1 [N25]; 981-999 of SEQ ID NO: 1 [C10]; 1051-1069 of SEQ ID NO: 1 [C15]; 1121-1139 of SEQ ID NO: 1 [C20]; and 1275-1296 of SEQ ID NO: 1 [C31], or a conservative variant or immunoreactive fragment thereof, wherein the presence of antibodies immunoreactive with said two or more amino acid sequences indicates immunoresistance to botulinum toxin therapy.

17. The method of claim 1, 6 or 10, wherein said botulinum toxin therapy is BoNT/A therapy.

18. A method of preventing or reducing immunoresistance to botulinum toxin therapy in an individual, comprising administering to said individual a tolerizing agent and two or more amino acid sequences selected from the group: 785-803 of SEQ ID NO: 1 [N25]; 981-999 of SEQ ID NO: 1 [C10]; 1051-1069 of SEQ ID NO: 1 [C15]; 1121-1139 of SEQ ID NO: 1 [C20]; and 1275-1296 of SEQ ID NO: 1 [C31], or a conservative variant or an immunoreactive fragment thereof, thereby preventing or reducing immunoresistance to botulinum toxin therapy.

30. The method of claim 18, 22 or 26, wherein said botulinum toxin therapy is BoNT/A therapy.

31. A method of vaccinating an individual against botulinum toxin, comprising administering to said individual a vaccine comprising an adjuvant and two or more amino acid sequences selected from the group 785-803 of SEQ ID NO: 1 [N25]; 981-999 of SEQ ID NO: 1 [C10]; 1051-1069 of SEQ ID NO: 1 [C15]; 1121-1139 of SEQ ID NO: 1 [C20]; and 1275-1296 of SEQ ID NO: 1 [C31], or a conservative variant or an immunoreactive fragment thereof, thereby producing an immune response to said botulinum toxin in said individual.

44. A method of removing botulinum toxin blocking antibodies from a patient, comprising the steps of (a) removing blood from a patient; (b) contacting said blood, or an antibody-containing component thereof with two or more amino acid sequences selected from the group 785-803 of SEQ ID NO: 1 [N25]; 981-999 of SEQ ID NO: 1 [C10]; 1051-1069 of SEQ ID NO: 1 [C15]; 1121-1139 of SEQ ID NO: 1 [C20]; and 1275-1296 of SEQ ID NO: 1 [C31], or a conservative variant or an immunoreactive fragment thereof, under conditions suitable for forming a complex of each of said amino acid sequences and anti-botulinum toxin antibody; and (c) removing said complex from said blood or antibody-containing component thereof.

47. The method of claim 44, 45 or 46, comprising selectively removing IgG botulinum toxin blocking antibodies from said patient.

48. A method of predicting or determining immunoresistance to botulinum toxin therapy in an individual, comprising the steps of: (a) determining the level of IgG antibodies immunoreactive with said botulinum toxin in said individual; and (b) comparing said level of IgG antibodies to a control level of IgG antibodies, wherein an increase in said level of IgG antibodies in said individual as compared to said control level indicates immunoresistance to said botulinum toxin therapy.

51. The method of claim 48, wherein said control level of IgG antibodies is determined in an individual who has not been treated with botulinum toxin therapy.

52. The method of claim 48, wherein said control level of IgG antibodies is determined in an individual who is responsive to said botulinum toxin therapy.

53. The method of claim 48, wherein said botulinum toxin therapy is BoNT/A therapy.

54. A method of predicting or determining immunoresistance to botulinum toxin therapy in an individual, comprising determining the presence or absence in said individual of antibodies immunoreactive with a BoNT/A peptide having a length of at most 60 amino acids and comprising an amino acid sequence selected from the group:

7 445-471 of SEQ ID NO: 1, 487-513 of SEQ ID NO: 1, 515-541 of SEQ ID NO: 1, 529-555 of SEQ ID NO: 1, 543-569 of SEQ ID NO: 1, 557-583 of SEQ ID NO: 1, 585-611 of SEQ ID NO: 1, 599-625 of SEQ ID NO: 1, 655-681 of SEQ ID NO: 1, 669-695 of SEQ ID NO: 1, 683-709 of SEQ ID NO: 1, 711-737 of SEQ ID NO: 1, 739-765 of SEQ ID NO: 1, 767-793 of SEQ ID NO: 1, 781-807 of SEQ ID NO: 1, 809-835 of SEQ ID NO: 1, 823-849 of SEQ ID NO: 1, and 837-863 of SEQ ID NO: 1,

or a conservative variant or immunoreactive fragment thereof, wherein the presence of antibodies immunoreactive with said peptide indicates immunoresistance to botulinum toxin therapy, and with the proviso that said BoNT/A peptide is not SEQ ID NO:2.

66. The method of claim 64, wherein said Hc peptide comprises an amino acid sequence selected from the group:

10 amino acids 939-957 of SEQ ID NO: 1 amino acids 953-971 of SEQ ID NO: 1 amino acids 967-985 of SEQ ID NO: 1 amino acids 981-999 of SEQ ID NO: 1 amino acids 995-1013 of SEQ ID NO: 1 amino acids 1009-1027 of SEQ ID NO: 1 amino acids 1023-1041 of SEQ ID NO: 1 amino acids 1037-1055 of SEQ ID NO: 1 amino acids 1051-1069 of SEQ ID NO: 1 amino acids 1065-1083 of SEQ ID NO: 1 amino acids 1079-1097 of SEQ ID NO: 1 amino acids 1093-1111 of SEQ ID NO: 1 amino acids 1107-1125 of SEQ ID NO: 1 amino acids 1121-1139 of SEQ ID NO: 1 amino acids 1135-1153 of SEQ ID NO: 1 amino acids 1149-1167 of SEQ ID NO: 1 amino acids 1163-1181 of SEQ ID NO: 1 amino acids 1177-1195 of SEQ ID NO: 1 amino acids 1191-1209 of SEQ ID NO: 1 amino acids 1205-1223 of SEQ ID NO: 1 amino acids 1219-1237 of SEQ ID NO: 1 amino acids 1233-1251 of SEQ ID NO: 1 amino acids 1247-1265 of SEQ ID NO: 1 amino acids 1261-1279 of SEQ ID NO: 1, and amino acids 1275-1296 of SEQ ID NO: 1,

or an immunoreactive fragment thereof.

73. The method of claim 54, wherein said botulinum toxin therapy is BoNT/A therapy.

74. A method of preventing or reducing immunoresistance to botulinum toxin therapy in an individual, comprising administering to said individual a tolerogizing agent and a BoNT/A peptide, said peptide having a length of at most 60 amino acids and comprising an amino acid sequence selected from the group: 445-471 of SEQ ID NO:1, 487-513 of SEQ ID NO:1, 515-541 of SEQ ID NO:1, 529-555 of SEQ ID NO:1, 543-569 of SEQ ID NO:1, 557-583 of SEQ ID NO:1, 585-611 of SEQ ID NO:1, 599-625 of SEQ ID NO:1, 655-681 of SEQ ID NO:1, 669-695 of SEQ ID NO:1, 683-709 of SEQ ID NO:1, 711-737 of SEQ ID NO:1, 739-765 of SEQ ID NO:1, 767-793 of SEQ ID NO:1, 781-807 of SEQ ID NO:1, 809-835 of SEQ ID NO:1, 823-849 of SEQ ID NO:1, and 837-863 of SEQ ID NO:1, or a conservative variant or immunoreactive fragment thereof, thereby preventing or reducing immunoresistance to

botulinum toxin therapy, with the proviso that said BoNT/A peptide is not SEQ ID NO:2.

82. The method of claim 74, wherein said tolerogizing agent and BoNT/A peptide are administered prior to said individual receiving botulinum toxin therapy.

83. The method of claim 82, wherein said individual is at increased risk for immunoresistance to botulinum toxin therapy.

84. A method of vaccinating an individual against botulinum toxin, comprising administering to said individual a vaccine comprising an adjuvant and a BoNT/A peptide, said peptide having a length of at most 60 amino acids and comprising an amino acid sequence selected from the group: 445-471 of SEQ ID NO:1, 487-513 of SEQ ID NO:1, 515-541 of SEQ ID NO:1, 529-555 of SEQ ID NO:1, 543-569 of SEQ ID NO:1, 557-583 of SEQ ID NO:1, 585-611 of SEQ ID NO:1, 599-625 of SEQ ID NO:1, 655-681 of SEQ ID NO:1, 669-695 of SEQ ID NO:1, 683-709 of SEQ ID NO:1, 711-737 of SEQ ID NO:1, 739-765 of SEQ ID NO:1, 767-793 of SEQ ID NO:1, 781-807 of SEQ ID NO:1, 809-835 of SEQ ID NO:1, 823-849 of SEQ ID NO:1, and 837-863 of SEQ ID NO:1, or a conservative variant or immunoreactive fragment thereof, thereby producing an immune response to botulinum toxin in said individual, with the proviso that said BoNT/A peptide is not SEQ ID NO:2.